AWARD NUMBER: W81XWH-14-1-0236

TITLE: Hippocampal and Cognitive Function, Exercise, and Ovarian Cancer: A Pilot Study

PRINCIPAL INVESTIGATOR: Richard Sloan, PhD

CONTRACTING ORGANIZATION: Columbia University

New York, NY 10032-3702

REPORT DATE: August 2016

TYPE OF REPORT: Annual Report

PREPARED FOR: U.S. Army Medical Research and Materiel Command

Fort Detrick, Maryland 21702-5012

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### 13. SUPPLEMENTARY NOTES

n/a

### 14. ABSTRACT

In this application, we propose to address this problem by focusing primarily on a single post-chemotherapy complaint in a single cancer: problems with memory in patients with ovarian cancer. We focus on this problem for three reasons: 1) according to the Ovarian Cancer National Alliance, memory problems are among the most frequently cited by patients; 2) by narrowly targeting our inquiry, we will avoid the "noise" in the data associated with different treatment regimens and correspondingly different cognitive complaints and; 3) there are decades of neuroscience research, some of it from our group, indicating that memory impairment, such as the kind reported in the context of chemotherapy, is mediated primarily by a region of the brain called the hippocampus. In this application, we propose to investigate the possibility that standard chemotherapy regimen used to treat ovarian cancer leads to memory impairment because it arrests the normal processes of neurogenesis, the growth of new nerve cells, in this brain region. In addition, because a growing body of research studies shows that physical exercise leads to improvement in memory and learning and that exercise targets the same brain regions responsible for chemotherapy-induced memory problems, we propose to conduct a pilot study of an intervention to increase patients' physical activity to test whether this will slow this effect of chemotherapy on nerve cell growth in the hippocampus and subsequently offset memory decline.

# 15. SUBJECT TERMS

Nothing listed

1 votiling listed					
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"Hippocampal and Cognitive Function, Exercise, and Ovarian Cancer: A Pilot Study"

**Proposal Log Number OC130386** 

Award Number W81XWH-14-1-0236

**HRPO Log Number A-18125** 

**Reporting Period: 8/1/15 – 7/31/16** 

### 1 INTRODUCTION:

In this application, we propose to address this problem by focusing primarily on a single post-chemotherapy complaint in a single cancer: problems with memory in patients with ovarian cancer. We focus on this problem for three reasons: 1) according to the Ovarian Cancer National Alliance, memory problems are among the most frequently cited by patients; 2) by narrowly targeting our inquiry, we will avoid the "noise" in the data associated with different treatment regimens and correspondingly different cognitive complaints and; 3) there are decades of neuroscience research, some of it from our group, indicating that memory impairment, such as the kind reported in the context of chemotherapy, is mediated primarily by a region of the brain called the hippocampus. In this application, we propose to investigate the possibility that standard chemotherapy regimen used to treat ovarian cancer leads to memory impairment because it arrests the normal processes of neurogenesis, the growth of new nerve cells, in this brain region. In addition, because a growing body of research studies shows that physical exercise leads to improvement in memory and learning and that exercise targets the same brain regions responsible for chemotherapy-induced memory problems, we propose to conduct a pilot study of an intervention to increase patients' physical activity to test whether this will slow this effect of chemotherapy on nerve cell growth in the hippocampus and subsequently offset memory decline.

# 2 KEYWORDS:

Physical activity interventions, ovarian cancer treatment, chemotherapy-induced cognitive dysfunction

### **3 ACCOMPLISHMENTS:**

# **Major Goals/Objectives:**

- 1. Approval by CUMC Herbert Irving Comprehensive Cancer Center (HICCC) Protocol Review and Monitoring Committee
  - Date of Completion: 4/17/14 (100% completed)
- 1A. Approval of HICCC protocol materials by DoD HRPO; Respond to DoD HRPO questions 100% completed
- 2.Approval by New York State Psychiatric Institute (NYSPI) Institutional Review Board Date of Completion: 5/15/15 (100% completed)
- 2A. Approval of NYSPI IRB materials by DoD HRPO; Respond to DoD HRPO questions Date of Completion: June 1, 2015 (100% completed)
- 3. Training research assistant "coach" in patient recruitment, retention, delivery of walking intervention Date of Completion: July 1, 2015 (100% completed)
- 4. Training research assistant in administration and scoring of neuropsychology tests Date of Completion: July 1, 2015 (100% completed)
- 5. Training research assistant in image analysis
  - Date of Completion: July 1, 2015 (100% completed)
- 6.Recruiting 21 ovarian cancer patients 4-6 weeks post-surgery
  - July 31, 2016: 5% completed
- 7.Pre- and post-intervention neuropsychology and imaging data collection and scoring July 31, 2016: 5% completed
- 8. Delivering the interventions to the two treatment groups

August 1, 2016: 5% completed

9.Data analysis

July 31, 2016: 5% completed

10.Post-Analysis: Submission of findings to national meetings, proposal to DoD or NIH for definitive study

(if appropriate)

July 31, 2016: 0% completed

# What was accomplished under these goals?

This is a study to assess the feasibility of recruiting ovarian cancer patients in a pilot study to assess the impact of increasing physical activity on neuropsychological and brain imaging outcomes. The principle finding to date is that due to changing treatment protocols, it is extremely difficult to recruit patients into the study. To date, we have screened 883 post-operative patients and have recruited only 1. With the exception of 4 patients, all others failed to meet inclusion/exclusion criteria. Of the 4 who met criteria, only one agreed to participate.

# What opportunities for training and professional development has the project provided?

Nothing to report

# How were the results disseminated to communities of interest?

Nothing to report

# What do you plan to do during the next reporting period to accomplish the goals?

We have expanded our potential pool of patients to those with endometrial and uterine cancers in the hope that we will be able to recruit more patients.

### **IMPACT:**

# What was the impact on the development of the principal discipline(s) of the project?

Nothing to report

# What was the impact on other disciplines?

Nothing to report

# What was the impact on technology transfer?

Nothing to report

# What was the impact on society beyond science and technology?

Nothing to report

# **CHANGES/PROBLEMS:**

# Changes in approach and reasons for change

To increase recruitment, we have opened enrollment to endometrial and uterine cancer patients.

# Actual or anticipated problems or delays and actions or plans to resolve them

As this is a feasibility study, we have encountered significant problems with recruitment, due in large part to changes in treatment protocols.

# Changes that had a significant impact on expenditures

Nothing to report

# Significant changes in use or care of human subjects, vertebrate animals, biohazards, and/or select agents

IRB approval to recruit endometrial and uterine cancer patients: January 16, 2016 IRB approval to recruit patients receiving both chemotherapy and radiotherapy: May 27, 2016

# Significant changes in use or care of human subjects

IRB approval to recruit endometrial and uterine cancer patients: January 16, 2016 IRB approval to recruit patients receiving both chemotherapy and radiotherapy: May 27, 2016

Significant changes in use or care of vertebrate animals.

Nothing to report

Significant changes in use of biohazards and/or select agents

Nothing to report

# **PRODUCTS:**

Publications, conference papers, and presentations:

Journal publications.

Nothing to report

Books or other non-periodical, one-time publications.

Nothing to report

Other publications, conference papers, and presentations.

Nothing to report

Website(s) or other Internet site(s)

Nothing to report

**Technologies or techniques** 

Nothing to report

Inventions, patent applications, and/or licenses

Nothing to report

**Other Products** 

Nothing to report

# PARTICIPANTS & OTHER COLLABORATING ORGANIZATIONS:

# What individuals have worked on the project?

Name:	Richard Sloan, PhD
Project Role:	Principal Investigator
Researcher Identifier (e.g. ORCID ID):	n/a
Nearest person month worked:	1
Contribution to Project:	Dr. Sloan is responsible for all aspects of the study including recruitment of patients, oversight of the activity-increasing program, and data analysis.
Funding Support:	This project.
Name:	Scott Small, MD
Project Role:	Co-Investigator
Researcher Identifier (e.g. ORCID ID):	n/a
Nearest person month worked:	1
Contribution to Project	Dr. Small supervises all aspects of the imaging within the
Contribution to Project:	proposed project and provides some supervision for the half-time Research Assistant in scoring the imaging data.
Funding Support:	This project.

Name:	Adam Brickman, PhD
Project Role:	Co-Investigator
Researcher Identifier (e.g. ORCID ID):	n/a
Nearest person month worked:	1
Contribution to Project:	Dr. Brickman will oversee the neuropsychological evaluation of participants in the study, including training study personnel on task administration, scoring, quality assurance, and data entry.
Funding Support:	This project.
Name:	Jason Wright, MD
Project Role:	Co-Investigator
Researcher Identifier (e.g. ORCID  ID):	n/a
Nearest person month worked:	I
Contribution to Project:	Dr. Wright will be responsible for patient recruitment as well as medical oversight for patients recruited to the study.
Funding Support:	This project.
Name:	Jose Henriquez-Rivera
Project Role:	Research Assistant
Researcher Identifier (e.g. ORCID ID):	n/a
Nearest person month worked:	12
Contribution to Project:	Mr. Henriquez-Rivera is the study "coach" and his responsibilities include patient recruitment, retention, delivery of walking intervention.
Funding Support:	This project.
Name:	Brianna Last
Project Role:	Research Assistant
Researcher Identifier (e.g. ORCID ID):	n/a
Nearest person month worked:	2
Contribution to Project:	Administering and scoring neuropsychological tests under Dr. Brickman's supervision.
Funding Support:	This project.
Name:	Christiane Hale
Project Role:	Research Worker
Researcher Identifier (e.g. ORCID ID):	n/a
Nearest person month worked:	1
Contribution to Project:	Administering and scoring neuropsychological tests under Dr. Brickman's supervision.

Funding Support:	This project.

# Has there been a change in the active other support of the PD/PI(s) or senior/key personnel since the last reporting period?

Yes, other support attached.

# What other organizations were involved as partners?

Nothing to report

# **SPECIAL REPORTING REQUIREMENTS:**

n/a

# **APPENDICIES**

n/a

#### Richard Sloan

Dr. Sloan has an appointment at the New York State Psychiatric Institute/Research Foundation for Mental Hygiene and Columbia University. This is detailed in a Dual Appointment Agreement between the entities and precludes the possibility of NYSPI/RFMH supporting effort supported by Columbia University and vice versa. Therefore, each project listed below indicates effort at each institution and also Total Professional Effort ('TPE') encompassing the multiple appointments.

### **Columbia University Active**

5R01HL128310-02 (Edmondson) 7/15/2015 - 3/31/2019 0.60 CM NIH/NHLBI TPE 0.49 CM

Test of a new theory to explain excess risk in cardiac PTSD

The goal of the proposed research is to identify targets for new interventions to reduce the doubled cardiac event recurrence and mortality risk faced by the 1 in 8 survivors of non-ST elevation myocardial infarction and unstable angina who develop PTSD secondary to their life-threatening cardiac event.

Role: Co-Investigator

 2P01AG003949-32 (Lipton)
 9/1/2017 - 5/31/2021
 1.80 CM

 Albert Einstein College of Medicine (NIH)
 TPE 1.47 CM

Einstein Aging Study

To identify risk factors and protective factors that influence both normative cognitive aging and the earliest onset of Alzheimer's Disease.

Role: Subcontract - Principal Investigator

Columbia CTSA Irving Institute (Picard) 9/1/2016 - 3/31/2017 0.12 CM Columbia CTSA Irving Institute TPE 0.10 CM

Stress Reactivity in Mitochondrial Disease: Preliminary Investigation of Physiological, Neural, and Epigenetic Mechanisms

This Collaborative and Multidisciplinary Pilot Research (CaMPR) Phase II award supports the establishment of the clinical and laboratory infrastructure for the MiSBIE (Mitochondrial Stress, Brain Imaging, and Epigenetics) study.

Role: Co-Investigator

5P01AG02166-10 (WISC) (Ryff) 7/1/2012 - 6/30/2017 1.01 CM NIH (WISC) TPE 1.20 CM

Integrative Pathways to Health & Fitness

The overall objective of MIDUS is to investigate the role of behavioral, psychological, and social factors in accounting for age-related variations in health and illness

Role: Subcontract - Principal Investigator

1U19AG051426-01A1 (WISC) (Ryff) 7/25/2016 - 5/31/2021 0.60 CM NIH (WISC) TPE 0.49 CM

Integrative Pathways to Health & Illness

The overall objective of MIDUS is to study health, broadly defined, as an integrated biopsychosocial process that unfolds across the decades of adult life.

Role: Subcontract - Principal Investigator

W81XWH-14-1-0236 (OC130386) (NCE) (Sloan) 8/1/2014 - 7/31/2017 0.25 CM
DOD TPE 0.30 CM

Hippocampal and Cognitive Function, Exercise and Ovarian Cancer: A Pilot Study

We propose to examine whether carboplatin and paclitaxel-based chemotherapy for ovarian cancer induces cognitive dysfunction and reduced DG CBV and to test whether an individually based walking intervention may attenuate this chemotherapy-induced cognitive and DG dysfunction.

Role: Principal Investigator

5P50AG008702-27 (Small) 6/1/2015 - 5/31/2020 0.12 CM NIH TPE 0.10 CM

Alzheimer's Disease Research Center

This project supports a wide spectrum of research on Alzheimer's disease.

Role: Co-Investigator

5 R01 AG035015-05 (NCE) (Small/Sloan) 9/15/2010 - 5/31/2017 1.01 CM NIH/NIA TPE 1.20 CM

Exercise, Age-Related Memory Decline and Hippocampal Function

The goal of this study is to conduct a randomized controlled trial of the effects of aerobic training and cognitive decline and to investigate the role of the hippocampus in mediating this effect.

Role: Multi-Principal Investigator

5 R01 AG033546-05 (NCE) (Stern/Sloan) 9/15/2010 - 5/31/2017 1.52 CM NIH/NIA TPE 1.80 CM

Exercise, Aging and Cognition: Effect and Mechanisms

The goal of this study is to extend the investigation of the beneficial effects of aerobic exercise to younger individuals, aged 25-40 and 50-65.

Role: Multi-Principal Investigator

#### **Columbia University Pending**

NIH (Chang) 7/1/2017 - 6/31/22 0.60 CM NHLBI/NIH TPE 0.49 CM

Test Of An Alternative ER Management Strategy For Tia Patients; An Explanatory And Guideline-Relevant Rct To Reduce PTSD After Tia And Minor Stroke This study will evaluate an outpatient treatment strategy for the management of transient ischemic attack (TIA) patients who present to the emergency department (ED).

Role: Co-Investigator

NIH (Cohen) 4/1/2017 - 3/31/2020 0.60 CM NIH TPE 0.49 CM

Does abdominal fat modify the effect of exercise on bone remodeling?

This proposal aims to establish, in a controlled study, to what extent exercise determines bone turnover based on serum markers. This marker data is required to plan biopsy studies to investigate effects of exercise on re-modeling at the tissue level. We hypothesize that low level of physical activity is an important and modi-fiable contributor to the low bone formation phenotype of abdominal obesity in premenopausal wom-en. We further hypothesize that increased exercise will ameliorate the bone formation defect and, ulti-mately, the bone quality defect found specifically in states of abdominal obesity.

Role: Co-Investigator

NIH (Diaz) 7/1/2017 - 6/30/2022 0.60 CM NIH TPE 0.49 CM

Breaking up prolonged sitting: a factorial trial to evaluate the efficacy of multiple components of a sitting interruption intervention for improving This study is designed to investigate the effects of a sitting interruption intervention in real world conditions over an extended period.

Role: Co-Investigator

NIH (Kimhy) 4/1/2017 - 3/31/2021 0.30 CM NIH TPE 0.24 CM

Individuals with schizophrenia (SZ) display substantial cognitive deficits across multiple domains. These deficits have been identified as major determinants of poor functioning and disability, representing a serious public health concern and an important target for interventions. At present, available pharmacological and cognitive-remediation treatments offer only minimal to limited benefits to ameliorate these deficits. Thus, there remains an urgent need to identify novel treatments for cognitive deficits in people with SZ.

Improving Cognition via Exercise in Schizophrenia

Role: Co-Investigator

PR160029 (Picard) 10/1/2017 - 9/30/2020 0.65 CM
Department of Defense TPE 0.53 CM

Stress Reactivity in Mitochondrial Disease: Physiological, Neural, and Epigenetic Mechanisms

This project will test the hypothesis that mitochondrial dysfunction in patients with the m.3243A>G mutation exhibit abnormal neural connectivity within the brain, which mediates exaggerated neuroendocrine, cardiovascular, and inflammatory responses to psychological stress, and abnormal epigenetic regulation of gene expression.

Role: Co-Investigator

1R01OD024671-01 (Picard) 9/1/2017 - 8/31/2022 1.20 CM NIH Director's Office TPE 0.98 CM

Mitochondrial and Neural Mechanisms Linking Stress to Disease

The goal of this transdisciplinary proposal is to define new principles of stress regulation in humans, and to explore novel sub-cellular mechanisms for mind-body processes.

Role: Co-Investigator

NIH (Sloan/Small) 4/1/2017 - 3/31/2022 3.00 CM NIH TPE 2.45 CM

Cocoa Flavanols, Systemic Inflammation, and Dentate Gyrus Function in Aging Adults

We propose to test this mechanism in a study in which healthy older adults will receive cocoa flavanols or placebo daily for a 12- week period, with preand post-intervention MRIs to assess the function of this brain region and blood draws. to measure key inflammatory markers.

Role: Principal Investigator

Department of Defense (Sun) 1/1/2017 - 6/30/2018 0.48 CM Department of Defense TPE 0.39 CM

 $Structured \& Regular \ Physical \ Activity \ in \ Children \ with \ Congenital \ Heart \ Diseases \ Can \ Enhance \ Neurocognitive \ Development$ 

The purpose of this Discovery Award application is to plan and complete all necessary preparations for a future randomized controlled trial (RCT) that will test the hypothesis: "Regular and structured physical activities in young children with congenital heart diseases will improve their neurodevelopmental outcome".

Role: Co-Investigator

### **OTHER SUPPORT**

**Active** 

W81XWH-14-1-0236 (Sloan)

08/01/2014 - 7/31/2017 (NCE)

0.30 calendar

Hippocampal and Cognitive Function, Exercise and Ovarian Cancer: A Pilot Study

In this application, we propose to investigate the possibility that standard chemotherapy regimen used to treat ovarian cancer leads to memory impairment because it arrests the normal processes of neurogenesis, the growth of new nerve cells, in this brain region.

Role on Project: Co-Investigator

R01AG016495 (Au)

09/15/2012 - 06/30/2017

0.12 calendar

NIH

MRI, Genetic and Cognitive Precursors of AD and Dementia

The goals of this study are to identify the most effective method and criteria for diagnosis of mild cognitive impairment (MCI) that are best predictive of incident Alzheimer's disease. Current efforts to treat AD have been ineffective because intervention occurs too late in the insidious process. Increased accuracy of diagnosis at the preclinical stage may be more successful for treatment and prevention efforts.

Role: Co-Investigator

1R01NS076837-01A1 (Cosentino, MPI)

09/21/2012 - 07/31/2017

0.12 calendar

NIH/NINDS

Examination of the Earliest Symptoms and Biomarkers of FTLD MAPT Carriers

This project examines the earliest clinical symptoms of FTLD in carriers of a tau mutation.

Role on Project: Co-investigator

P50AG008702 (Small/Brickman-Project Leader)

06/01/2015 - 05/31/2020

1.35 calendar

NIH/NIA

Alzheimer's Disease Research Center

This project supports a wide spectrum of research on Alzheimer's disease.

Role on Project: Project PI

1U01AG051412-01(Schupf, Lott, Silverman) 10/01/2015 - 09/30/2020

1.80 calendar

NIA

Biomarkers of Alzheimer's Disease in adults with Down Syndrome

The goal of this interinstitutional/collaborative study focuses on a longitudinal and multidisciplinary determination of key biomarkers that are likely to define this progression, including levels and rates of change in blood based biomarkers such as □-amyloid peptides, protein and lipid profiles, and measures of amyloid and tau concentration in cerebrospinal fluid, neuroimaging-based changes and genetic polymorphisms Role on Project: Co-investigator and Site PI for Massachusetts General Hospital

5R01CA186080-02 (Greenlee)

04/15/2015 - 03/31/2020

0.15 calendar

NCI

Cook for Your Life: Maintaining dietary change among breast cancer survivors

Conduct a 2x2 factorial randomized controlled trial to test the effects of a hands-on nutritional education curriculum on changing dietary behaviors among a diverse population of English- and Spanish-speaking breast cancer survivors who have completed treatment.

Role on Project: Co-Investigator

1RF1AG051556-01(Brickman, Luchsinger, Moreno) 09/30/2015 - 08/31/2020

0.90 calendar

Interdisciplinary Research to Understand the Interplay of Diabetes, Cerebrovascular disease and Alzheimer's Disease

### **OTHER SUPPORT**

To conduct studies in humans and mice relating diabetes to Alzheimer's and vascular mechanisms. Role on Project: Co-PI

R01AG049810 (Bondi)

03/15/2016-02/28/2021

0.90 calendar

NIH

Re-visiting Methods for MCI Diagnosis to Improve Biomarker and Trial Findings

Subcontract site is to provide collaborative support for the aims and goals of the UCSD-based research project Role on Project: Co-I

1R01AG050440-01A1 (Luchsinger)

09/01/2015 - 05/31/2020

0.90 calendar

NIH/NIA

Diabetes Status and Brain Amyloid in Middle Aged Hispanics

The main goal of this proposal is to study whether diabetes status (type 2 diabetes [referred to as diabetes] and pre-diabetes, compared with normal glucose tolerance [NGT]), is related to increased amyloid ß (Aß) deposition in the brain, one of the culprits of Alzheimer's disease (AD), in a community sample of 150 middle aged Hispanics with a mean age of 63 years.

Role on Project: Co-Investigator

RF1AG054023 (Mayeux)

08/01/2016 - 06/30/2021

0.30 calendar

NIH

Genetic Epidemiology of Cerebrovascular Factors in Alzheimer's Disease

The overall goal of this project is to test hypotheses concerning how genetic variants, cardiovascular risk factors, and cerebrovascular disease predispose to Late Onset Alzheimer's disease and whether these relationships differ by ethnic group.

Role on Project: Co-Investigator

RF1AG054070 (Manly/Brickman)

09/01/2016 - 07/31/2021

3.00

calendar NIH/NIA

Offspring Study of Mechanisms for Racial Disparities in Alzheimer's Disease

The overall aim of this study is to identify biological and sociocultural mechanisms of racial/ethnic disparities in cognitive function among middle-aged people with and without a parent with Alzheimer's Disease.

Role on Project: Co-PI

2R56AG034189-06A1 (Brickman)

09/01/2016 - 08/31/2021

1.80 calendar

NIH/NIA

White Matter Hyperintentsities in Aging and Dementia

This project will examine the degree to which small vessel cerebrovascular contributes independently or interactively to the development and clinical expression of AD across racial and ethnic groups. It will provide novel mechanistic insight into the disease and help identify new targets for intervention.

R00AG47963 (Zahodne)

09/01/2016 - 08/31/2019

0.36 calendar

NIH/NIA

Psychosocial protective factors in cognitive and brain aging

Characterizing potential intervention targets to reduce age-related cognitive morbidity in diverse elders is of critical importance to the U.S. aging population. This project aims to determine which positive psychosocial factors (1) buffer the impact of brain pathology on cognition and (2) protect against cognitive decline in older adults of different racial/ethnic backgrounds and with different cognitive abilities.

# <u>Pending</u>

### **OTHER SUPPORT**

PAR-16-365 (Luchsinger)

10/01/2016 - 09/30/2021

0.60 calendar

NIH

Mediterranean Diet in Alzheimer's dementia Prevention (MEDAP)

To conduct a randomized trial comparing the Mediterranean diet vs. health education among 200 persons with prodromal Alzheimer's disease to obtain preliminary data on feasibility and efficacy in preventing cognitive decline

Role on Project: Co-I

PAR-15-349 (Luchsinger)

10/01/2016 - 09/30/2021

1.20 calendar

NIH

Ethnic Disparities in Brain Amyloid and Cognition in Middle Age

To compare disparities in brain amyloid and cognition among late middle aged Hispanics, Non-Hispanic Blacks, and Non-Hispanic Whites in Northern Manhattan.

Role on Project: Co-I

3RF1AG051556-01S2 (Brickman, Luchsinger, Moreno) 10/01/2016 - 08/31/2020

0.30 calendar

NIA

Interdisciplinary Research to Understand the Interplay of Diabetes, Cerebrovascular disease and Alzheimer's Disease

This revision is to add Tau PET imaging to human studies in order to complement animal studies that are examining tau neuropathology.

Role on Project: Co-PI

3R01AG050440-02S1 (Luchsinger)

10/01/2016 - 08/31/2020

0.30 calendar

NIH/NIA

Diabetes Status and Brain Amyloid in Middle Aged Hispanics

This revision proposes to add Tau PET imaging to the ongoing amyloid PET imaging

Role on Project: Co-I

PAR-15-349 (Zahodne)

12/01/2016 - 08/31/2021

1.20

calendar NIH/NIA

Resilience Mechanisms Underlying Racial/Ethnic Disparities in Alzheimer's Disease

Identifying modifiable factors that contribute to racial/ethnic disparities in Alzheimer's disease (AD) is of critical importance to the increasingly diverse U.S. aging population. Using repeat MRI and cognitive assessments across three time points, this study examines how psychological and other resources that differ across race/ethnicity promote resilience at multiple points in the AD pathogenic pathway.

Role on Project: Corresponding PI

R21AG056940 (Brickman)

07/01/2017-06/30/2019

1.44 calendar

NIH/NIA

Cerebral autoregulation, Alzheimer's biomarkers, and white matter hyperintensities

This project will examine the interrelationship among cerebral autoregulation, small vessel cerebrovascular disease, AD cerebrospinal fluid biomarkers, and cognitive functioning in older adults. It will provide novel mechanistic insight into the disease and help identify new targets for intervention or disease prevention. Role on Project: PI

Parent R01 (Rutherford)

04/01/2017 - 03/31/2022

1.20 calendar

NIH

Trajectories of Healthy Aging: Linking Neuropathology, Emotion, and Late-Life Depression

Role on Project: Co-Investigator

### **OTHER SUPPORT**

R01AG055299 (Luchsinger)

07/01/2017 - 06/31/2022

1.20 calendar

NIH

Are there ethnic differences in brain amyloid and tau in the seventh decade of life?

To compare the presence of brain amyloid and tau in-vivo between Blacks, Hispanics, and Whites in the seventh decade of life

Role on Project: Co-Investigator

W81XWH-14-1-0236 (Sloan)

08/01/2017 - 7/31/2020

0.30 calendar

DoD

Hippocampal and Cognitive Function, Exercise and Ovarian Cancer: A Pilot Study

In this application, we propose to investigate the possibility that standard chemotherapy regimen used to treat ovarian cancer leads to memory impairment because it arrests the normal processes of neurogenesis, the growth of new nerve cells, in this brain region.

Role on Project: Co-Investigator

### Overlap

# For New and Competing Applications (PHS 398) – DO NOT SUBMIT UNLESS REQUESTED For Non-competing Progress Reports (PHS 2590) – Submit only Active Support for Key Personnel

### PHS 398/2590 OTHER SUPPORT

# Jason D. Wright

**ACTIVE** 

NCI R01 CA169121-01A1 (Wright, PI)

1/1/13-12/31/17

3.0 calendar

**National Cancer Institute** 

The Influence of Hospital Variability on the Management of Cancer-Associated Complications

To evaluate common complications from cancer therapy using administrative data.

W81XWH-14-1-0236 (Sloan, PI)

8/1/14-7/31/16

0.3 calendar

Department of Defense

Hippocampal and Cognitive Function, Exercise, and Ovarian Cancer: A Pilot Study

The major goals of this award are to determine the cognitive effects of exercise in ovarian cancer survivors.

SAC 160066 (Hershman, PI)

5/1/16-4/30/19

1.2 calendar

Susan G. Komen Foundation

Comorbidity, Toxicity, and Breast Caner Survival Among Women On and Off Clinical Trials.

The goal of this project is to examine the effect of comorbidities on survival for women with breast cancer treated on cooperative group trials and in general practice.

### **OVERLAP**

There is no scientific or budgetary overlap on any current project.

# **PENDING**

**American Cancer Society** 

1.2 calendar

Oral Anticancer Drugs: Expense, Access, Adherence, and Safety

To evaluate barriers to use, adherence, cost and comparative effectiveness of tyrosine kinase inhibitors in patients with solid tumors.

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# SCOTT A. SMALL, M.D. Other Support

# **Active**

1R01AG035015 (Small/Sloan)

09/15/10-05/31/17 NCE

0.56 CM NIH/

NIA

# Exercise, Age-Related Memory Decline, and Hippocampal Function

The goal of this proposal is to conduct a randomized controlled trial of the effects of aerobic training on cognitive decline and to investigate the role of the hippocampus in mediating this effect.

U01 AG016976 (Small)

07/01/98-06/30/19

0.10 CM

University of Washington Subcontract

NACC

Goals are to coordinate activities of Alzheimer's Centers by working with and supplying data to national center.

R01 AG042317 (Abeliovich)

07/01/12-06/30/17

0.26 CM

NIH/NIA

### Human induced neuronal stem cell models of familial Alzheimer's disease

To determine the mechanism for altered endosomal trafficking in Familial Presenilin mutant Alzheimer's disease models.

W81XWH-14-1-0236 (Sloan)

8/1/14 - 7/31/17

0.3 CM

DoD

# Hippocampal and Cognitive Function, Exercise and Ovarian Cancer: A Pilot Study

In this application, we propose to investigate the possibility that standard chemotherapy regimen used to treat ovarian cancer leads to memory impairment because it arrests the normal processes of neurogenesis, the growth of new nerve cells, in this brain region.

9527 (Abelovich) 09/1/14 - 8/31/17

1.68 CM

Michael J. Fox Foundation

# Targeting retromer dysfunction: a convergent mechanism in familial and sporadic PD

The overarching goals of the present proposal are A.) To further detail mechanisms by which LRRK2 and other PD-related genes regulate vesicular trafficking in cells, and identify potential therapeutic targets and B.) To develop in vitro and in vivo models and tools, including drug-like reagents, for further studies regarding potential therapeutic targets such as VPS35.

Agreement #46560 (Small)

01/14/2013-12/31/2016

0.6 CM

**Anonymous** 

*In vivo validation of the retromer complex as a key component of Alzheimer's disease etilogy* Establish a link between retromer dysfunction and tau toxicity.

1R01MH093398 (Small)

09/22/11-06/30/22

0.60 CM

NIMH

# Longitudinal Imaging of Patients at Clinical Risk for Psychosis

In this proposal we will use a variant of functional brain imaging that can detect disease-associated dysfunction in small regions of the brain and apply this to patients at clinical risk for psychosis who are followed prospectively for clinical and brain imaging outcomes. The main project goal is to definitively test the hypothesis of hippocampal hyperfunction as a pathogenic driver in schizophrenia and related disorders.

P50 AG08702 (Small)

09/29/89-05/31/20

1.16 CM

NIH/NIA

# Alzheimer's disease Research Center

This project supports a wide spectrum of research on Alzheimer's disease.

# **Pending**

# <u>n/a</u>

# Overlap

There is no scientific overlap between any of the grants listed above and the application under consideration. If the pending grants are funded, efforts will be adjusted accordingly.